

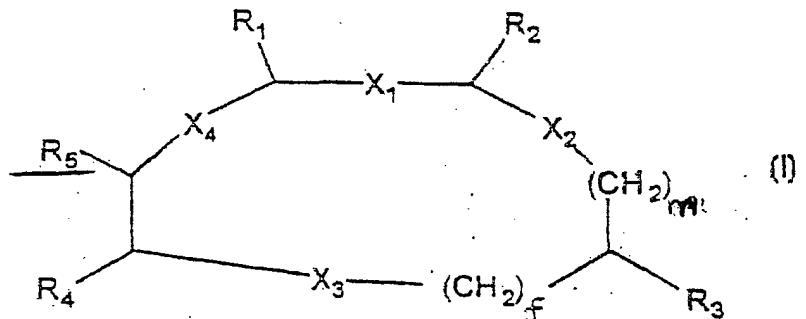
IN THE CLAIMS

21. (Currently Amended) Monocyclic compounds of formula (I)

wherein:

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15 X_1, X_2, X_3, X_4 are the same or different, and are selected from the group consisting of -CONR-, -NR₂CO-, -CH₂-NR-, and -NR-CH₂- where R is selected from the group consisting of H, C₁₋₃ alkyl, and benzyl;

f and m are the same or different, and is are a number selected from the group consisting of 0, 1 and 2;

20 R_1 and R_2 , are the same or different, and represent: -(CH₂)_rAr where r is 0, 1 or 2 and Ar is an aromatic group selected from the group consisting of benzene, naphthalene, thiophene, benzothiophene, pyridine, quinoline, indole, furan, benzofuran, thiazole, benzothiazole, imidazole, benzimidazole, optionally substituted with up to 2 substituents selected from the group consisting of C₁₋₃ alkyl, C₁₋₃ haloalkyl, C₁₋₃ alkyloxy, C₂₋₄ amino-alkyloxy, halogens, OH, NH₂, CN, and NR₆R₇, where R₆ and R₇, same or different, are H or C₁₋₃ alkyl,

25 R_3 is -(CH₂)_rAr where r is 0, 1 or 2 and Ar is an aromatic group selected from the group consisting of benzene, naphthalene, thiophene, benzothiophene, pyridine, quinoline, indole, furan, benzofuran, thiazole, benzothiazole, imidazole, and benzimidazole,

optionally substituted with up to 2 groups selected from the group consisting of C₁₋₃ alkyl, C₁₋₃ haloalkyl, C₁₋₃ alkyloxy, amino-alkyloxy, halogens, OH, NH₂, and NR₆R₇, where R₆ and R₇, same or different, are H or C₁₋₃ alkyl,

R₄ is-NR₈R₉/; -N(R₁₁)CO(CH₂)_hR₁₂; or -COR₁₃; where R₅ is H; where R₈ is H or C₁₋₃ alkyl; h is 0,

5 1, 2 or 3: and R₉ is selected from the group consisting of methanesulfonyl, tosyl, tetrahydropyranyl, tetrahydrothiopyranyl optionally mono or di-substituted by oxygen on the S atom, piperidyl, optionally substituted on the N-atom by a C₁₋₃ alkyl, C₁₋₃ acyl, aminosulfonyl, or methanesulfonyl; or a group-(CH₂)_gR₁₀ where g is 1,2, or 3 and R₁₀ is selected from the group consisting of morpholine, furan and CN;

10 or R₈ and R₉ together with the N atom to which they are linked form a piperazine optionally substituted at the other N atom substituted by a C₁₋₃ alkyl, C₁₋₃ acyl or methanesulfonyl; -N(R₁₁)CO(CH₂)_hR₁₂ where R₁₁ is H or C₁₋₃ alkyl; h is 0, 1, 2 or 3; and R₁₂ is selected from the group consisting of morpholine, pyrrolidine optionally substituted with a hydroxy or hydroxymethyl, piperidine optionally substituted with a 4-hydroxy/ or 4-carboxyamido or 4-aminosulfonyl group;

15 piperazine optionally substituted on the N-atom by 4-aminosulfonyl, C₁₋₃ alkyl, triazole, tetrazole, 5-mercpto-tetrazole, furan, thiophene, thiomorpholine, optionally mono or di-oxygenated on the S atom, and amino-cyclohexane cyclohexan-1-yl-amino optionally substituted by a hydroxy group; -COR₁₃ wherein R₁₃ is a member selected from the group consisting of morpholine and piperazine optionally substituted by a C₂₋₆ alkyl containing one or more hydroxy groups;

20 their enantiomers and mixtures thereof, their diastereoisomers, and their pharmaceutically acceptable salts.

22. (Currently Amended)

Compound according to Claim 21 wherein:

f is 1

m is 0

X_1, X_2, X_3, X_4 , are the same or different and are a member selected from the group consisting of -CONR- and -NRCO-,

5 where R is H or methyl,

R_1 and R_2 are the same or different, are:

- CH_2Ar wherein Ar is an aromatic group selected from the group consisting of benzene, pyridine, indole, optionally substituted with up to two substituents selected from the group consisting of C_{1-3} alkyl, C_{1-3} haloalkyl, C_{1-3} alkyloxy, C_{2-4} amino alkyloxy, halogens, OH, NH_2 , CN, and NR_6R_7 ,
- 10 where R_6 and R_7 , same or different, and are H or C_{1-3} alkyl;
- R_3 is $-CH_2Ar_4$ wherein Ar_4 is an aromatic group selected from the group consisting of alpha naphthyl, beta naphthyl, phenyl, phenyl substituted with up to two substituents selected from the group consisting of C_{1-3} alkyl, C_{1-3} haloalkyl, C_{1-3} alkyloxy, halogens, OH, and NH_2 .

23. (Currently Amended) Compounds according to Claim 22 wherein:

- 15 - X_1, X_2, X_3, X_4 are -CONH-,
 - R_1 is indol-3-yl-methyl
 - R_2 is phenyl-methyl optionally substituted with up to two substituents selected from the group consisting of chlorine, fluorine, CF_3 , OH and CN ; or are is selected from the group consisting of 3-pyridyl-methyl and 4-pyridyl-methyl;
- 20 - R_3 is benzyl.

24. (Previously Added) Compounds according to claim 23 wherein:

R_4 is a group NR_8R_9 wherein:

25 R_8 is H or methyl;

R_9 selected from the group consisting of 4-tetrahydropyranyl, 4-tetrahydrothiopyranyl, 1-oxo-tetrahydrothiopyran-4-yl, 1,1-dioxo-tetrahydrothiopyran-4-yl, N-methyl-4-piperidinyl, N-methanesulfonyl-4-piperidinyl, and N-aminosulfonyl-4-piperidinyl,
or R_8 and R_9 together with the N atom to which they are linked represent N-methyl-piperazinyl,
5 N-acetyl-piperazinyl or N-methanesulfonyl-piperazinyl.

25. (Previously Amended) Compounds according to Claim 24 represented by:

- i) cyclo{Suc[1-(R)-(4-tetrahydropyranyl)amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}
- 10 ii) cyclo{Suc[1-(S)-(4-tetrahydropyranyl)amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}
- iii) cyclo{Suc[1-(R)-(1-methyl-piperidin-4-yl)amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}
- iv) cyclo{Suc[1-(R)-(4-tetrahydrothiopyranyl)amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}
- 15 v) cyclo{Suc[1-(R)-(1-oxo-tetrahydrothiopyran-4-yl)amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}
- vi) cyclo{Suc[1-(R)-(1,1-dioxo-tetrahydrothiopyran-4-yl)amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}
- 15 vii) cyclo{Suc[1-(R)-N-methyl-N-(4-tetrahydropyranyl)amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}
- viii) cyclo{Suc[1-(R)-(4-tetrahydropyranyl)amino]-Trp-Tyr-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}
- 20 ix) cyclo{Suc[1-(R)-(4-tetrahydropyranyl)amino]-Trp-Phe(4-F)-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}
- x) cyclo{Suc[1-(R)-(4-tetrahydropyranyl)amino]-Trp-Phe(3,5-F)-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}
- xi) cyclo{Suc[1-(R)-(4-tetrahydropyranyl)amino]-Trp-Phe(4-CN)-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}
- 25 xii) cyclo{Suc[1-(R)-(4-tetrahydropyranyl)amino]-Trp-Phe(4-CF₃)-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

-CH₂NH]}{

xiii) cyclo{Suc[1-(R)-(4-tetrahydropyanyl)amino]-Trp-Ala(4-pyridyl)-[(R)-NH-CH(CH₂-C₆H₅)

-CH₂NH]}{

xiv) cyclo{Suc[1-(R)-(4-tetrahydropyanyl)amino]-Trp-Ala(3-pyridyl)-[(R)-NH-CH(CH₂-C₆H₅)

5 -CH₂NH]}{

xv) cyclo{Suc[1-(R)-(1-methylsulfonyl-piperidin-4-yl)amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)

-CH₂NH]}{

xvi) cyclo{Suc[1-(R)-(1-aminosulfonyl-piperidin-4-yl)amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)

-CH₂NH]}{

10 xvii) cyclo{Suc[1-(R)-4-methyl-piperazin-1-yl]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

xviii) cyclo{Suc[1-(R)-4-acetyl-piperazin-1-yl]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{ or

xix) cyclo{Suc[1-(R)-4-methylsulfonyl-piperazin-1-yl]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{.

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26. (Previously Amended) Compounds according to Claim 23 wherein:

R₄ represents a group NR₈R₉, where R₈ is H and R₉ is methanesulfonyl, tosyl or a group

-(CH₂)_gR₁₀, wherein g is 1 or 2 and R₁₀ is morpholine, furan, or CN.

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27. (Previously Amended) Compounds according to claim 26 represented by:

xx) cyclo{Suc[1-(S)-methylsulfonylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

xxi) cyclo{Suc[1-(R)-methylsulfonylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

xxii) cyclo{Suc[1-(S)-(4-methylphenyl)sulfonylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

xxiii) cyclo{Suc[1-(R)-(4-methylphenyl)sulfonylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

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xxiv) cyclo{Suc[1-(S)-2-(4-morpholino)ethylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

xxv) cyclo{Suc[1-(R)-2-(4-morpholino)ethylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

xxvi) cyclo{Suc[1-(R)-(2-furyl)methylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]} or

xxvii) cyclo{Suc[1-(R)-cianomethylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}.

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28. (Previously Added) Compounds according to claim 23 wherein:

R₄ is a group - N(R₁₁)CO(CH₂)_h-R₁₂ wherein R₁₁ is H, h is 0 or 1, and R₁₂ is selected from the group consisting of 1-tetrazolyl, 5-mercaptop-tetrazol-1-yl, 1-triazolyl, furanyl, thiophenyl, morpholine, 4-10 hydroxy-piperidine, 4-carboxyamido-piperidine, 3-hydroxy-pyrrolidine, 2-hydroxymethylpyrrolidine, 4-methyl-piperazine, 4-aminosulfonyl-piperazine, 1-oxo-thiomorpholine, and 4-hydroxy-cyclohexan-1-yl-amino.

29. (Previously Added) Compounds according to Claim 28 represented by:

15 xxviii) cyclo{Suc[1-(R)-2-(4-morpholino)acetylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

xxix) cyclo{Suc[1-(S)-2-(4-morpholino)acetylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

xxx) cyclo{Suc[1-(S)-2-(tetrazol-1-yl)acetylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

xxxi) cyclo{Suc[1-(R)-2-(tetrazol-1-yl)acetylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

20 xxxii) cyclo{Suc[1-(S)-2-(5-mercaptop-tetrazol-1-yl)acetylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

xxxiii) cyclo{Suc[1-(R)-2-([1,2,4]triazol-1-yl)acetylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

xxxiv) cyclo{Suc[1-(R)-2-(furanyl)carbonylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

25 xxxv) cyclo{Suc[1-(R)-2-(thiophen-3-yl)acetylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

xxxvi) cyclo{Suc[1-(R)-(4-morpholino)carbonylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

xxxvii) cyclo{Suc[1-(R)-2-(4-hydroxy-piperidin-1-yl)acetylamino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}

-C₆H₅)-CH₂NH]}{

xxxviii) cyclo{Suc[1-(R)-2-(4-aminocarbonyl-piperidin-1-yl)acetyl amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

xxxix) cyclo{Suc[1-(R)-2-(3-hydroxy-pyrrolidin-1-yl)acetyl amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

5 xli) cyclo{Suc[1-(R)-2-(2-(S)-hydroxymethyl-pyrrolidin-1-yl)acetyl amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

xlii) cyclo{Suc[1-(R)-2-(4-methyl-piperazin-1-yl)acetyl amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

10 xlvi) cyclo{Suc[1-(R)-2-(4-methyl-piperazin-1-yl)carbonyl amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

xliii) cyclo{Suc[1-(R)-2-(4-aminosulfonyl-piperazin-1-yl)acetyl amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{

15 xliv) cyclo{Suc[1-(R)-2-(1-oxo-thiomorpholin-4-yl)acetyl amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{ or

xlvi) cyclo{Suc[1-(R)-2-(*trans*-4-hydroxy-cyclohexan-1-yl-amino)acetyl amino]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{.

20 30. (Previously Amended) Compounds according to Claim 23 wherein:

R₄ represents a group COR₁₃ wherein R₁₃ is morpholine.

31. (Previously Amended) Compounds according to claim 30 represented by:

25 xlvi) cyclo{Suc[1-(4-morpholino)carbonyl]-Trp-Phe-[(R)-NH-CH(CH₂-C₆H₅)-CH₂NH]}{.

32. (Previously Added) Pharmaceutical compositions containing as active principle compounds of general formula (I) according to Claim 21 in combination with pharmaceutically acceptable carriers or excipients.

33. (Previously Added) A method for the treatment of the bronchospastic component of 5 asthma, cough, pulmonary irritation, intestinal spasms or local spasms of bladder, ureters during cystitis, kidney infections and colics wherein amounts of 0.1 to 10mg/kg body weight of an active principle represented by compounds of formula (I) according to Claim 21 are administered to the patient.